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AMENDMENTS TO THE CLAIMS

The following listing of claims will replace all prior versions and listings of claims in the application.

Listing of claims:

Claim 1 (currently amended).

A compound selected from those of formula

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$$(R_2)_{m} \xrightarrow{A} (Z_1)_{n} \xrightarrow{Z} X_1 \xrightarrow{N} W$$

in which:

R₁ represents <u>hvdrogen</u>, a group selected from:

- -hydrogen, amino,
- (C₁-C₆)alkyl, (C₂-C₆)alkenyl, (C₃-C₆)alkynyl, mono(C₁-C₆)alkylamino(C₁-C₆)alkyl, di(C₁-C₆)alkylamino(C₁-C₆)alkyl, aryl, aryl(C₁-C₆)alkyl, heterocycle, and 3- to 6 membered cycloalkyl(C₁-C₆)alkyl, those groups being unsubstituted or substituted with one-or-more groups, which may be identical or different, selected from amino, (C₁-C₆)alkyl, eyano, halo(C₁-C₆)alkyl, C(=O)OR₄, OR₄ and SR₄, in which R₄-represents hydrogen or (C₁-C₆)alkyl,

W represents an oxygen atom, a sulphur atom, or a group =N-R', in which R' represents (C_1-C_6) alkyl, hydroxyl, or cyano,

X₁ and X₃ represent, independently of each other, a group -C-R₆ in which R₆ represents a group selected from hydrogen, (C₁-C₆)alkyl, amino,

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mono(C_1 - C_6)alkylamino, di(C_1 - C_6)alkylamino, hydroxyl, (C_1 - C_6)alkoxy, and halogen;

X₂ is nitrogen;

Y represents a group selected from oxygen atom, sulphur atom, -NH, and - $N(C_1-C_6)$ alkyl,

Z represents:

- · an oxygen atom, a sulphur atom,
- or a group -NR₇ in which R₇ represents a group selected from hydrogen, (C₁-C₆)alkyl, aryl(C₁-C₆)alkyl, cycloalkyl, aryl, and heteroaryl, and
- when Y is an oxygen atom, a sulphur atom, or a group $-N(C_1-C_6)$ alkyl, Z optionally represents a carbon atom which is unsubstituted or substituted with a (C_1-C_6) alkyl, an aryl, an aryl (C_1-C_6) alkyl, an aromatic or non-aromatic heterocycle or a cycloalkyl,

n is an integer from 1 to 8 inclusive,

 Z_1 represents $-CR_8R_9$ wherein R_8 and R_9 , independently of each other, represent a group selected from hydrogen, (C_1-C_6) alkyl, halo (C_1-C_6) alkyl, halogen, amino, OR_4 , SR_4 or $C(=O)OR_4$ in which R_4 represents a hydrogen or (C_1-C_6) alkyl, and

- when n is greater than or equal to 2, the hydrocarbon chain Z₁ optionally contains one or more multiple bonds,
- and/or one of the carbon atoms in the hydrocarbon chain Z₁ may be replaced with an oxygen atom, a sulphur atom which is unsubstituted or substituted with one or two oxygen atoms, or a nitrogen atom which is unsubstituted or substituted with a (C₁-C₆)alkyl,
- and when one of the carbon atoms in the hydrocarbon chain Z_1 is replaced with a sulphur atom which is unsubstituted or substituted with one or two oxygen

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atoms, then the group -C(=Y)-Z- optionally may be absent in the general formula (I),

A represents an a group selected from:

- aromatic or non-aromatic, 5- or 6-membered monocycle comprising from 0
 to 4 heteroatoms selected from nitrogen, oxygen and sulphur, and
- bicycle, composed of two aromatic or non-aromatic, 5 or 6 membered rings, which may be identical or different, comprising from 0 to 4 heteroatoms selected from nitrogen, oxygen and sulphur,

m is an integer from 0 to 7 inclusive,

the group(s) R_2 , which may be identical or different, is (are) selected from (C_1-C_6) alkyl, halogen, -CN, NO₂, SCF₃, -CF₃, -OCF₃, -NR₁₀R₁₁, -OR₁₀, -SR₁₀, SOR₁₀, -SO₂R₁₀, -(CH₂)_kSO₂NR₁₀R₁₁, -X₅(CH₂)_kC(=O)OR₁₀, -(CH₂)_kC(=O)NR₁₀R₁₁, -(CH₂)_kC(=O)NR₁₀R₁₁, and -X₄-R₁₂ in which:

- X₅ represents a group selected from oxygen, sulphur optionally substituted by one or two oxygen atoms, and nitrogen substituted by hydrogen or (C₁-C₆)alkyl,
 - k is an integer from 0 to 3 inclusive,
- R₁₀ and R₁₁, which may be identical or different, are selected from hydrogen and (C₁-C₆)alkyl,
- X₄ represents a group selected from single bond, -CH₂-, oxygen atom, sulphur atom optionally substituted by one or two oxygen atoms, and nitrogen atom substituted by hydrogen atom or (C₁-C₆)alkyl group,

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• R₁₂ represents an aromatic or non-aromatic, heterocyclic or non-heterocyclic, 5- or 6-membered ring which is unsubstituted or substituted with one or more groups, which may be identical or different, selected from (C₁-C₆)alkyl, halogen, hydroxyl and amino, and when the ring is heterocyclic, it comprises from 1 to 4 heteroatoms selected from nitrogen, oxygen and sulphur,

Ra represents a group selected from:

- hydrogen,
- (C₁-C₆)alkyl, (C₃-C₆)alkenyl, (C₃-C₆)alkynyl, these groups being unsubstituted or substituted with one or more groups, which may be identical or different, selected from amino, cyano, halo(C₁-C₆)alkyl, cycloalkyl, C(=O)NR₁₀R₁₄, C(=O)OR₁₀, OR₁₀, and SR₁₀, in which R₁₀ and R₁₁, which may be identical or different, represent hydrogen or (C₁-C₆)alkyl,
 - and the group of formula:

- ✓ in which p is an integer from 0 to 8 inclusive,
- ✓ Z₂ represents -CR₁₃R₁₄ wherein R₁₃ and R₁₄, independently of each other, represent a group selected from hydrogen, (C₁-C₆)alkyl, phenyl, halo(C₁-C₆)alkyl, halogen, amino, OR₄, SR₄ and -C(=O)OR₄ in which R₄ represents hydrogen or (C₁-C₆)alkyl, and
 - when p is greater than or equal to 2, the hydrocarbon chain Z₂ optionally contains one or more multiple bonds,
 - and/or one of the carbon atoms in the hydrocarbon chain Z_2 may be replaced with an oxygen atom, a sulphur atom which is unsubstituted or substituted with one or two oxygen atoms, a nitrogen atom which is unsubstituted or substituted with a (C_1-C_6) alkyl, or a carbonyl group,

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- ✓ B represents a group selected from:
 - an aromatic or non-aromatic 5- or 6-membered monocycle comprising from 0 to 4 heteroatoms selected from nitrogen, oxygen and sulphur, and

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- a bicycle, composed of two aromatic or non-aromatic, 5- or 6-membered rings, which may be identical or different, comprising from 0 to 4 heteroatoms selected from nitrogen, oxygen and sulphur,
- ✓ q is an integer from 0 to 7 inclusive,
- ✓ the group(s) R₅, which may be identical or different, is (are) selected from CF₃, OCF₃, $-(CH_2)_kNR_{15}R_{16}$ (C_1-C_6) alkyl, halogen, CN, NO_2 , $-N(R_{15})C(=O)R_{16}, -N(R_{15})C(=O)OR_{16}, -N(R_{15})SO_2\hat{R}_{16}, -N(SO_2R_{15})_2, -OR_{15},$ -(CH₂)_kSO₂NR₁₅R₁₆, $-SO_2-N(R_{15})-(CH_2)_{k2}-NR_{16}R_{17}$ $-S(O)_{k1}R_{15}$ $-(CH_2)_kC(=O)OR_{15}$, $-C(=O)O-(CH_2)_{k2}-NR_{15}R_{16}$, $-X_7(CH_2)_kC(=O)OR_{15}$, $-X_7(CH_2)_kC(=O)NR_{15}R_{16}$ $-C(=O)O-(CH_2)_{k2}-C(=O)OR_{18}$ -(CH₂)_kC(=O)NR₁₅R₁₆, -R₁₉-C(=O)OR₁₅, -X₆-R₂₀, and -C(=O)-R₂₁-NR₁₅R₁₆ in which:
 - X₇ represents a group selected from oxygen atom, sulphur atom optionally substituted by one or two oxygen atoms, and nitrogen atom substituted by a hydrogen atom or a (C₁-C₆)alkyl group,
 - k is an integer from 0 to 3 inclusive,
 - k1 is an integer from 0 to 2 inclusive,
 - k2 is an integer from 1 to 4 inclusive,

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- R₁₅, R₁₆ and R₁₇, which may be identical or different, are selected from hydrogen and (C₁-C₆)alkyl,
- R₁₈ represents a group selected from (C₁-C₆)alkyl, -R₂₁-NR₁₅R₁₆, -R₂₁-NR₁₅-C(=O)-R₂₁-NR₁₆R₁₇, and -C(=O)O-R₂₁-NR₁₅R₁₆ in which R₂₁ represents a linear or branched (C₁-C₆)alkylene group, and R₁₅, R₁₆ and R₁₇ are as defined hereinbefore,
- R₁₉ represents a (C₃-C₆)cycloalkyl group,
- X_6 represents a group selected from single bond, -CH₂-, oxygen atom, sulphur atom optionally substituted by one or two oxygen atoms, and nitrogen atom substituted by hydrogen atom or (C₁-C₆)alkyl group.
- R₂₀ represents an aromatic or non-aromatic, heterocyclic or non-heterocyclic, 5- or 6-membered ring, which is unsubstituted or substituted with one or more groups, which may be identical or different, selected from (C₁-C₆)alkyl, halogen, hydroxyl, oxo, cyano, tetrazole, amino, and -C(=O)OR₄ wherein R₄ represents hydrogen or (C₁-C₆)alkyl, and, when the ring is heterocyclic, it comprises from 1 to 4 heteroatoms selected from nitrogen, oxygen and sulphur,

with the provise that when X₁ represents a nitrogen atom, X₂ cannot represent a carbon atom substituted with a methyl group or with NH CH₂; optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claim 2 (canceled).

Claim 3 (currently amended). A compound of formula (I) according to Claim 1 characterized in that:

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n is an integer from 1 to 6 inclusive,

Z₁ represents -CR₈R₉ wherein R₈ represents a hydrogen atom and R₉ represents a hydrogen atom or a methyl group, and

- when n is greater than or equal to 2, the hydrocarbon chain Z_1 optionally contains a double bond,
- or, one of the carbon atoms in the hydrocarbon chain Z_1 may be replaced with an oxygen atom, or a sulphur atom which is unsubstituted or substituted with one or two oxygens,

A represents a group selected from phenyl, pyridyl, thienyl, imidazolyl, furyl, and piperidyl, 1,3-benzodioxolyl, benzodioxinyl, benzothienyl, benzofuryl, benzofuryl, benzofuryl, 2,1,3-benzothiadiazolyl, and indolyl,

m is an integer from 0 to 7 inclusive,

the group(s) R_2 , which may be identical or different, is (are) selected from (C_1-C_6) alkyl, halogen, -CN, $-CF_3$, $-OCF_3$, $-NR_{10}R_{11}$, $-OR_{10}$, $-SR_{10}$, $-SO_2R_{10}$, $-(CH_2)_kSO_2NR_{10}R_{11}$, $-X_5(CH_2)_kC(=O)OR_{10}$, $-(CH_2)_kC(=O)OR_{10}$,

- $-X_5(CH_2)_kC(=0)NR_{10}R_{11}$, $-(CH_2)_kC(=0)NR_{10}R_{11}$, and $-X_4-R_{12}$ in which:
- ✓ X₅ represents O, S or NH,
- ✓ k is an integer from 0 to 3 inclusive,
- \checkmark R₁₀ and R₁₁, identical or different, are selected from hydrogen and (C₁-C₆)alkyl,
- ✓ X₄ represents -CH₂-, or an oxygen atom,
- ✓ R₁₂ represents a phenyl group which is unsubstituted or substituted with one
 or more groups, which may be identical or different, selected from
 (C₁-C6)alkyl, halogen, hydroxyl and amino,

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optionally, the racernic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claim 4 (currently amended). A compound of formula (I) according to Claim 1 characterized in that:

R₃ represents hydrogen, (C₁-C₆)alleyl-or the group of formula:

$$(R_s)_q$$
 B $(Z_2)_p$

- in which p is an integer from 0 to 3 inclusive,
- Z₂ represents -CR₁₃R₁₄ wherein R₁₃ and R₁₄, independently of each other, represent a group selected from hydrogen, methyl, or phenyl, and
 - when p is greater than or equal to 2, the hydrocarbon chain Z₂ optionally contains one double bond,
 - or one of the carbon atoms in the hydrocarbon chain Z₂ may be replaced with an oxygen atom, a sulphur atom which is unsubstituted or substituted with one or two oxygen atoms, a nitrogen atom which is unsubstituted or substituted with a (C₁-C₆)alkyl, or a carbonyl group,
- B represents a group selected from phenyl, pyridyl, thienyl, imidazolyl, furyl, 1,3-benzodioxolyl, benzodioxinyl, benzothienyl, benzofuryl, 2,1,3-benzothiadiazolyl, benzofurazanyl, naphthyl, and indolyl,
- q is an integer from 0 to 3 inclusive,
- the group(s) R₅, which may be identical or different, is (are) selected from (C₁-C₆)alkyl, halogen, CN, NO₂, CF₃, OCF₃, -(CH₂)_kNR₁₅R₁₆,
 -N(R₁₅)C(=O)R₁₆, -(R₁₅)C(=O)OR₁₆, -N(R₁₅)SO₂R₁₆, -N(SO₂R₁₅)₂, -OR₁₅,
 -S(O)_{k1}R₁₅, -SO₂-N(R₁₅)-(CH₂)_{k2}-NR₁₆R₁₇, -(CH₂)_kSO₂NR₁₅R₁₆,

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- $-X_7(CH_2)_kC(=O)OR_{15}, -(CH_2)_kC(=O)OR_{15}, -C(=O)O-(CH_2)_{k2}-NR_{15}R_{16}, \\ -X_7(CH_2)_kC(=O)NR_{15}R_{16}, \text{ and } -(CH_2)_kC(=O)NR_{15}R_{16} \text{ in which :}$
- X₇ is \$, 0 or NH,
- k is an integer from 0 to 3 inclusive,
- k1 is an integer from 0 to 2 inclusive,
- k2 is an integer from 1 to 4 inclusive,
- R₁₅, R₁₆ and R₁₇, which may be identical or different, are selected from hydrogen and (C₁-C₆)alkyl,

optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claims 5 to 8 (canceled).

Claim 9 (previously presented). A compound of formula (I) according to Claim 1 characterized in that:

W represents an oxygen atom,
Y represents an oxygen atom,
Z represents a NH group,
Z₁ represents a methylene group,
and n is equal to one,

optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claims 10 and 11 (canceled).

Claim 12 (previously presented). A compound of formula (I) according to Claim 1 characterized in that: X_1 and X_3 represent each a -CH group, and

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optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claim 13 (currently amended). A compound of formula (I) according to Claim 1 characterized in that:

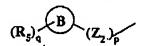
A represents a group selected from phenyl, and pyridyl, 1,3 benzedioxolyl, and benzefurazanyl,

m is equal to 0 or 1,

and R_2 represents a group selected from (C_1-C_6) alkoxy, hydroxy, halogen, and (C_1-C_6) thioalkoxy,

optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claim 14 (previously presented). A compound of formula (I) according to Claim 1 characterized in that R₃ represents a group of formula:



in which:

p is equal to 1,

Z₂ represents a methylene group,

B represents a group selected from phenyl, pyridyl, 1,3-benzodioxolyl, and benzofurazanyl,

q is an integer from 0 to 2 inclusive,

and R₅ represent(s) a group selected from halogen, CN, -(CH₂)_kNR₁₅R₁₆,

- $-S(O)_{k1}R_{15}, -(CH_2)_kSO_2NR_{15}R_{16}, -(CH_2)_kC(=O)OR_{15}, -(CH_2)_kC(=O)NR_{15}R_{16}, \text{ and } -(CH_2)_kC(=O)R_{15}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}R_{16}, -(CH_2)_kC(=O)R_{16}R_{16}R_{16}, -(CH_2)_kC(=O)R$
- -X₆-R₂₀, in which:
- k is an integer from 0 to 1 inclusive,
- k1 is an integer from 0 to 2 inclusive,
- R₁₅ and R₁₆, which may be identical or different, are selected from hydrogen and (C₁-C₆)alkyl,

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- X₆ represents a bond,
- R₂₀ represents a 5-membered heterocyclic ring comprising from 3 to 4 heteroatoms selected from oxygen and nitrogen and optionally substituted with a methyl group or an oxo group.

optionally, the racemic forms thereof, isomers thereof, N-oxides thereof, and the pharmaceutically acceptable salts thereof.

Claim 15 (currently amended). A compound of formula (I) according to Claim 1, which is:

- _____1,3_Dimethyl_2,4-dioxo_1,2,3,4_tetrahydro-pyrido[3,4_d]pyrimidine_6-earboxylic_acid_(1,3_benzodioxol_5_ylmothyl)_antide,
- 3-Benzyl-1 methyl-2,4-dioxo-1,2,3,4-tetrahydro pyrido[3,4-d] pyrimidine 6-carboxylic acid (1,3-benzodioxol-5 ylmethyl) amide,
- Methyl 4-[6-(4-Methoxy-benzylcarbamoyl)-1-methyl-2,4-dioxo-1,4-dihydro-2H-pyrido[3,4-d]pyrimidin-3-ylmethyl]-benzoate,
- 4-[6-(4-Methoxy-benzylcarbamoyl)-1-methyl-2,4-dioxo-1,4-dihydro-2*H*-pyrido[3,4-*d*]pyrimidin-3-ylmethyl]-benzoic acid,
- 4-[6-(3-Methoxy-benzylcarbamoyl)-1-methyl-2,4-dioxo-1,4-dihydro-2*H*-pyrido[3,4-*d*]pyrimidin-3-ylmethyl]-benzoic acid, and
- 3-(4-Cyano-benzyl)-1-methyl-2,4-dioxo-1,2,3,4-tetrahydro-pyrido[3,4-d]pyrimidine-6-carboxylic acid 4-methoxy-benzylamide.

Claims 16 to 28 (canceled).

Claim 29 (currently amended). A pharmaceutical composition comprising a compound according to any one of Claims 1, 3, 4, 8, 9, 11-15 Claims 1, 3, 4, 9, or 11-15 inclusive and a pharmaceutically acceptable excipient.

Claims 30 and 36 (canceled).

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Claim 37 (currently amended). A method for treating a disease or complaint involving a therapy by inhibition of MMP-13 characterized in that the disease or the complaint is arthritis, the method comprising the administration of an effective amount of a compound according to any one of Claims 1, 3, 4, 8, 9, 11 16 Claims 1, 3, 4, 9, or 11-15 inclusive to a patient having arthritis.

Claim 38 (currently amended). A method for treating according to Claim 37, charaotherized characterized in that the disease is osteoarthritis.

Claim 39 (currently amended). A method for treating according to Claim 37, charactherized characterized in that the disease is rheumatoid arthritis.